

\bigcup_{PhNION} A review of deep learning and radiomics approaches for pancreatic cancer diagnosis from medical imaging

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Purpose of review

Early and accurate diagnosis of pancreatic cancer is crucial for improving patient outcomes, and artificial intelligence (AI) algorithms have the potential to play a vital role in computer-aided diagnosis of pancreatic cancer. In this review, we aim to provide the latest and relevant advances in AI, specifically deep learning (DL) and radiomics approaches, for pancreatic cancer diagnosis using cross-sectional imaging examinations such as computed tomography (CT) and magnetic resonance imaging (MRI).

Recent findings

This review highlights the recent developments in DL techniques applied to medical imaging, including convolutional neural networks (CNNs), transformer-based models, and novel deep learning architectures that focus on multitype pancreatic lesions, multiorgan and multitumor segmentation, as well as incorporating auxiliary information. We also discuss advancements in radiomics, such as improved imaging feature extraction, optimized machine learning classifiers and integration with clinical data. Furthermore, we explore implementing AI-based clinical decision support systems for pancreatic cancer diagnosis using medical imaging in practical settings.

Summary

Deep learning and radiomics with medical imaging have demonstrated strong potential to improve diagnostic accuracy of pancreatic cancer, facilitate personalized treatment planning, and identify prognostic and predictive biomarkers. However, challenges remain in translating research findings into clinical practice. More studies are required focusing on refining these methods, addressing significant limitations, and developing integrative approaches for data analysis to further advance the field of pancreatic cancer diagnosis.

Keywords

computer-aided diagnosis, computed tomography, deep learning, medical imaging, magnetic resonance imaging, pancreatic cancer, radiomics

INTRODUCTION

Clinical motivation

Pancreatic cancer is one of the most lethal cancers with a poor prognosis and limited treatment options, with a 5-year relative survival rate of only 12% [\[1\].](#page-10-0) Over 90% of pancreatic cancers are exocrine tumors, for which the most common type is pancreatic ductal adenocarcinoma (PDAC), as illustrated in Fig. 1. There is no widely accepted screening for pancreatic cancer yet, and cross-sectional imaging is still the choice of noninvasive method that is widely available [\[2\]](#page-10-0).

Identifying early precursor changes in the pancreas has the potential to aid in the risk prediction of pancreatic cancer, as these precancerous alterations manifest as morphological and textural changes on abdominal imaging (CT and MRI) [\[3,4,42,43\].](#page-10-0) These changes include main pancreatic duct (MPD) stricture and upstream marked MPD dilatation [\[5\]](#page-10-0). This

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KEY POINTS

- Deep learning and radiomics show strong potential for improving pancreatic cancer diagnosis using computed tomography and magnetic resonance imaging.
- Recent advances in pancreatic cancer diagnosis combine radiomics methods with machine learning classifiers, while deep learning approaches employ diverse architectures for automated tumor detection, classification, and multiorgan segmentation, demonstrating significant progress in medical imaging analysis.
- Challenges that must be addressed include developing fusion models, establishing large public datasets, improving generalizability, and integrating diagnostic functions into computer-aided diagnosis systems.
- Future research should focus on refining methods, addressing limitations, and developing integrative approaches for data analysis.

observation provides the rationale for our research on computer-aided diagnosis (CAD) methods, specifically deep learning (DL) and radiomics, in identifying pancreatic cancer risk at an early stage. However, detecting these early changes has challenges such as:

- (1) variable size, shape and texture of the pancreatic lesions,
- (2) limited volume of medical datasets due to cost of examination and manual annotation,
- (3) variable imaging techniques collected from different institutions.

Despite these challenges, early detection is crucial, given the high mortality rate associated with pancreatic cancer. Therefore, it is essential to develop robust and reliable CAD systems for medical imaging to improve pancreatic cancer diagnosis.

Multiple imaging modalities are available for diagnosis of pancreas cancer: CT, MRI, endoscopic ultrasound (EUS), and transabdominal ultrasound. In this review, we primarily focus on CT and MRI as they are the most common noninvasive diagnostic imaging methods with certain superiorities available. Among CT and MRI, CT is the most utilized modality in cancer diagnosis due to its availability, high spatial resolution, and fast acquisition times. However, MRI offers superior soft tissue contrast resolution and provide more insight into tissue characteristics such as signal intensity on T1- or T2-weighted images, better visualization of ductal strictures, as well as advanced imaging, such as diffusion-weighted imaging (DWI) and perfusion

imaging. Figure 1 shows both MRI and corresponding EUS images of pancreatic cancer.

Cross-sectional imaging is the most commonly used modality for the diagnosis of pancreatic cancer. However, this practice has challenges, as tumors might be missed or can be confused with other benign pancreatic lesions or abnormalities. CAD systems have been developed over years to alleviate some of the difficulties in the current standard. For example, a CAD system can help identify small tumors in the pancreas region that may be missed or help predict patient outcomes after diagnosis of pancreatic cancer, surgical planning, or therapy response assessment in radiation oncology. Related to the role of artificial intelligence (AI) in pancreatic cancer diagnosis, we explore two prominent families of methods of medical image classification for pancreatic cancer that have garnered significant attention and demonstrated success: deep learning and radiomics. Gaining a comprehensive understanding of the current state of research in these areas is essential for enhancing diagnostic accuracy and improving patient outcomes. We discuss recent advances, challenges, and future perspectives too.

Development of radiomics and artificial **intelligence**

A typical workflow of a CAD system for pancreatic cancer diagnosis involves image preprocessing, region of interest (ROI) segmentation, and tumor classification, after which the results are presented to a clinician for further evaluation and treatment/ surgical planning or response assessment (Fig. 2). Radiomics and DL are essential in tumor segmentation, classification, and early diagnosis. Although radiomics is often considered a pre-DL era method of extracting hand-crafted features for auto-diagnosis, DL is regarded as a feature exploration method for the same tasks. DL can be used for tumor segmentation, crucial for diagnosis and prognostic purposes because the size, shape, and location of the tumor are critical in such tasks.

These advanced techniques (DL and radiomics) are becoming increasingly significant as they can detect subtle changes and aid in the early detection and characterization of pancreatic tumors. One may improve diagnostic accuracy in differentiating benign and malignant lesions by leveraging quantitative imaging biomarkers extracted from radiomics and high-level features from deep learning algorithms. However, challenges remain regarding image registration (alignment), data availability, clinical interpretability, robustness and generalization ability.

FIGURE 1. Pancreatic ductal adenocarcinoma (PDAC). (a) T1-weighted (T1W) MRI image with fat suppression (FS); (b) contrast-enhanced (CE) T1W MRI image during the early phase in same patient; (c) PDAC; (d) axial CE-T1W-FS image with PDAC in the uncinate process of the head; (e) axial CE-T1W-FS image with PDAC and liver metastases; (f) EUS image showing PDAC measuring 2.5 cm. The white arrows point to cancer. EUS, endoscopic ultrasound; PDAC, pancreatic ductal adenocarcinoma.

Evaluation criteria

Evaluation metrics are essential for providing clear, objective, and quantifiable ways to assess and compare research findings; therefore, herein, we enlist commonly used evaluation strategies in CAD systems for identifying weaknesses and strengths of available methods. Several widely used metrics from the confusion matrix and the receiver operating characteristic (ROC) curve are employed for determining the effectiveness and reliability (Fig. 3): sensitivity, specificity, accuracy, precision, recall, Fscore and area under the receiver operating characteristic curve (AUC). The key metrics are explained here:

(1) Sensitivity: Sensitivity, also known as the true positive rate, measures the proportion of confirmed cases that the CAD system correctly identifies as positive. It is a measure of the system's ability to detect cancerous lesions in medical images. High sensitivity implies proficiency at identifying pancreatic cancer cases, minimizing the chances of false-negative results.

- (2) Specificity: Specificity, or the true negative rate, reflects the system's ability to distinguish noncancerous tissue from cancers. High specificity indicates that the CAD system effectively avoids false-positive results, reducing the likelihood of unnecessary additional tests or treatments.
- (3) Accuracy: Sometimes, balanced accuracy is used to evaluate performance on an unbalanced dataset. Accuracy represents the proportion of all cases (both positive and negative) that the CAD system correctly classifies. In pancreatic cancer diagnosis, accuracy is an overall measure of the

FIGURE 2. A typical pancreatic cancer CAD system workflow, including two strategies, deep learning and radiomics. CAD, computer-aided diagnosis.

CAD system's performance in correctly distinguishing cancerous and noncancerous lesions.

(4) AUC: ROC curve is a graphical representation of a classifier's performance across various threshold settings. The curve plots the true positive rate (TPR $=$ sensitivity) against the false positive rate (FPR $= 1$ – specificity) at different thresholds. The AUC is a summary metric that quantifies the overall performance of the CAD system across all possible thresholds. As shown in Fig. 3, an AUC of 1 (line C-A-D) indicates perfect classification, while an AUC of 0.5 (dotted line C-D) suggests that the CAD system's performance is no better than random chance. A high AUC value implies that the CAD system effectively distinguishes between pancreatic cancer and noncancer cases across a range of threshold settings.

In conclusion, these evaluation metrics provide a comprehensive performance assessment, as each metric contributes unique insights into the classifier's ability to detect cancerous lesions and avoid false diagnoses.

Search criteria

The literature search keywords used to identify relevant articles include pancreatic cancer, computed tomography (CT), magnetic resonance imaging (MRI), pancreatic ductal adenocarcinoma (PDAC), intraductal papillary mucinous neoplasms (IPMN), pancreatic cysts, auto-diagnosis, CAD systems for pancreatic cancer, and a mixture of them. We used Google Scholar, PubMed, and Scopus to conduct a literature search and manually reviewed the reference lists of selected articles to identify additional relevant studies. Articles were included if they met the following criteria:

- (1) Published within the specified time window (November 2021–April 2023).
- (2) Focused on pancreatic cancer diagnosis using CT or MRI.

FIGURE 3. Common evaluation metrics. Left: confusion matrix; right: ROC curve. ROC, receiver operating characteristic.

- (3) Employed radiomics, deep learning, or a combination of both methods for image analysis.
- (4) Investigated pancreatic cancer types or predisposing conditions mentioned above.

RADIOMICS

Basics

Radiomics has been applied to pancreatic cancer imaging data to provide prognostic and predictive information that can guide clinical decision making. Radiomics is an advanced image analysis technique that systematically extracts quantitative features from images, known as manually crafted features [\[5\].](#page-10-0) These features capture detailed information about the morphology, texture, and intensity distribution within the pancreas. The radiomics workflow typically involves several steps, including image preprocessing, region of interest (ROI) segmentation, feature extraction, feature selection, and classification or prediction using machine learning algorithms (Fig. 4).

The primary advantage of radiomics lies in its ability to provide an extensive and objective assessment of pancreas heterogeneity, which can significantly enhance clinical decision-making. By leveraging high-dimensional feature spaces, radiomics enables the discovery of complex patterns and correlations that may not be readily apparent through traditional visual inspection by physicians (or radiologists). Consequently, radiomics has shown promise [\[6–8,44\]](#page-10-0) in improving diagnostic accuracy with an AUC of 0.7–0.8, depending on the specific dataset and implementation $[9^{\bullet},10^{\bullet} [9^{\bullet},10^{\bullet} [9^{\bullet},10^{\bullet} 12", 13… 14".$

Recent advances

Recent advances in radiomics methods for pancreatic cancer diagnosis tasks have demonstrated significant progress in medical imaging analysis, benefiting from a mature methodology pipeline. These studies often integrate clinical features, employ feature selection techniques, and compare the performance of various machine learning classifiers, as illustrated in detail in Table 1.

A common practice in these studies is to use feature selection methods to identify the most relevant radiomics features for tumor classification. Ma *et al*. [\[9](#page-10-0)**"**] and Flammia *et al.* [\[13](#page-10-0)**""**] both utilized the least absolute shrinkage and selection operator (LASSO) method to select features that were then integrated into their respective models. LASSO is a linear regression analysis method often used in statistics and machine learning as a regularization strategy to prevent overfitting. Moreover, LASSO offers greater interpretability than other complex machine learning and statistical methods, as it shrinks less important parameters to zero while explaining the regression results using the remaining coefficients. This feature makes LASSO particularly useful for identifying the most relevant radiomics features and building more transparent models. Another study by Ahmad *et al*. [\[10](#page-10-0)**"**] employed a naïve Bayes classifier after selecting features potentially predictive of PDAC. Naive Bayes classification is often used when features are considered conditionally independent given the output label. Incorporating clinical features is beneficial [\[9](#page-10-0)"], and researchers continue investigating more informative hand-crafted features $[13$ ^{H}].

In model development, various machine learning classifiers have been used in these studies [9",10"-12",13"",14"]. Support Vector Machines

FIGURE 4. A detailed radiomics-based diagnosis workflow.

Table 1. Recent advances in deep learning for pancreatic cancer diagnosis based on imaging. Ground truth diagnoses in the dataset are usually based on pathology data

AUC, area under the receiver operating characteristic curve; CAD, computer-aided diagnosis; CNN, convolutional neural network.

(SVM) and XGBoost have emerged as robust classifiers for such tasks $[11^\bullet,12^\bullet,14^\bullet]$, with most advances focusing on application aspects, such as the com-parison between CT and MRI [\[14](#page-10-0)"]. Experiments on diverse datasets from different regions have further validated the value of radiomics in pancreatic cancer diagnosis tasks [\[15\]](#page-10-0).

DEEP LEARNING

Basics

Deep learning has demonstrated remarkable potential in pancreatic cancer diagnosis, with numerous studies reporting promising results [\[16–20\]](#page-10-0). Widely utilized models include CNN-based architectures

such as UNet and its variations [\[21\]](#page-10-0). Transfer learning is often employed to adapt pretrained CNN models, including VGG [\[22\]](#page-10-0), Inception [\[23\]](#page-10-0), and ResNet [\[24\],](#page-10-0) to discern specific features in pancreatic cancer imaging. Despite its promise, deep learning faces challenges such as data scarcity, model interpretability, and generalizability. Several data augmentation techniques are frequently employed to address these issues. [\[16,25,26\]](#page-10-0), increasing the size and diversity of training datasets, thereby enhancing model performance on previously unseen images or domains.

Figure 5 illustrates a simplified diagram of commonly-employed deep learning approaches. To explain the differences among deep learning strategies: CNN is the most typical deep learning architecture, consisting of multiple convolutional layers and other network architecture elements, and is being successfully used for many image classifications, segmentation and detection tasks. Transformers, on the contrary, have recently gained momentum and have been shown to be state-ofthe-art, replacing CNN-based methods. The main premise behind the Transformers is to have a procedure called self-attention mechanism, allowing them one to weigh and consider different parts of

the images when making predictions. Lastly, generative adversarial network (GAN) based methods encompass encoder, generator, and discriminator components, functioning in two stages: training and validation. These are substantially different from CNNs and transformers in those GANs are often used to generate new data samples, while CNNs and Transformers are used for classification and detection tasks. One should note that GANs are generative models, and CNNs and Transformers are discriminative models. All these families of deep learning methods have been used in pancreatic cancer detection and analysis studies; therefore, it is essential to enlist their basic architecture and differences, as in Fig. 5.

Recent advances

Most research until today were conducted on CT images, with a few investigations involving MRI [\[33](#page-10-0)"] and EUS [\[27,28\]](#page-10-0). Several studies have developed end-to-end DL solutions for pancreatic tumor detection and classification. For instance, Park *et al*. [\[29](#page-10-0)**"**], Althobaiti *et al*. [\[30](#page-10-0)"], and Chen *et al*. [\[31](#page-10-0)""] employed different deep learning architectures, such as nnU-Net, Capsule Network, and CNNs, respectively, for

FIGURE 5. Deep learning architectures used for pancreatic cancer diagnosis. The diagram is organized into three main categories: CNN-based models, transformer-based models, and cycleGAN. GAN, generative adversarial network; CNN, convolutional neural network.

detecting and segmenting pancreatic tumors. These studies further show the potential of deep learning in automating the diagnosis process and reducing the workload for radiologists.

Despite further investigation into CNN-based models, recent advances in deep learning methods for pancreatic cancer diagnosis have led to the development of transformer-based models, which have demonstrated significant potential in various aspects of medical imaging. Transformer models are known for making robust encoders [\[37\]](#page-10-0), and vision transformers have been specifically applied for the classification of intraductal papillary mucosal neoplasms (IPMN) in MRI images $[33$ ^{H}].

In addition to using established deep learning architectures, some studies have proposed novel deep learning models. For instance, Vaiyapuri *et al*. [\[32](#page-10-0)**"**] introduced the IDLDMS-PTC, a system that combines MobileNet for feature extraction with an optimal autoencoder and multileader optimization for classification. Another study introduced a meta-information-aware dual-path transformer for pancreatic lesion classification and segmentation, highlighting the potential of incorporating auxiliary information to improve model performance [\[36](#page-10-0)["]].

Performance details are summarized in the following Table 2. It is worth noting that dataset bias could potentially impact performance. Nevertheless, these recent advancements highlight the growing potential of transformer-based deep learning techniques in the diagnosis and management of pancreatic cancer.

Notably, there has been a growing interest in developing deep learning algorithms that can recognize multiple tumor types or provide multiorgan and multitumor semantic segmentation. Chen *et al.'*s [\[35](#page-10-0)**"**] Unified Tumor Transformer (UniT) model exemplify this approach, detecting and diagnosing eight major cancers from CT scans. Furthermore, the FELIX Project presents a suite of deep learning algorithms designed to recognize pancreatic lesions, specifically PDAC $[34"$.

DISCUSSION

In summary, there are exciting recent advances in AI, specifically DL and radiomics approaches, for the diagnosis of pancreatic cancer using medical imaging modalities such as CT and MRI. Looking forward, there are multiple potential avenues for future research as well as a need to address the challenges in translating these findings into clinical practice.

Fusion models

Integrating DL and radiomics approaches could potentially lead to more powerful diagnostic tools for pancreatic cancer. Combining these methods may result in enhanced model performance, benefiting from the strengths of both techniques, as suggested by several studies [\[38–40\].](#page-10-0) Future research should investigate the development and application of fusion models in pancreatic cancer diagnosis using medical imaging.

Public large datasets and benchmarks

Most studies discussed in this review utilized internal datasets, which could introduce bias when comparing model performance. There is an urgent need to develop large volume public datasets and standardized benchmarks to address this issue. These resources would facilitate more accurate model comparisons and improve the generalizability of findings across different patient populations and institutions.

Generalizability across tumor types, domains, and organs

Future research should focus on developing models that can be used across different types of pancreatic neoplasia, such as PDAC and IPMN. Additionally, there is potential for creating models capable of generalizing across organs, which could improve healthcare practice by enabling the diagnosis of various tumor types using a single model. However, this presents challenges, as different organs have varying difficulty levels regarding segmentation and classification. For instance, abdominal segmentation, particularly for the pancreas, is more challenging than brain segmentation due to the complex shapes and structures involved. Developing automatic or self-adaptive models that can effectively address these challenges remains an open question.

Integration of diagnostic functions into computer-aided diagnosis systems

Recent research has begun to explore the integration of various diagnostic procedures, such as preprocessing, segmentation, and classification, into CAD systems for pancreatic cancer. This holistic approach could streamline the diagnostic process and improve the efficiency of medical imagingbased pancreatic cancer diagnosis. Future studies should continue investigating the development and implementation of such integrated CAD systems.

DL and radiomics with medical imaging have shown great promise for improving the diagnostic accuracy of pancreatic cancer, facilitating

Table 2. Recent advances in deep learning for pancreatic cancer diagnosis based on imaging. Ground truth diagnoses in the dataset are usually based on clinical manifestations and pathology data

CNN, convolutional neural network; PDAC, pancreatic ductal adenocarcinoma.

personalized treatment planning, and identifying prognostic and predictive biomarkers. However, challenges remain in translating these research findings into clinical practice. By addressing these challenges and focusing on the areas discussed above, future research can continue to advance the field of pancreatic cancer diagnosis using medical imaging.

CONCLUSION

In conclusion, the application of artificial intelligence techniques, specifically deep learning and radiomics, for the diagnosis of pancreatic cancer (especially at the early stage) using cross-sectional imaging modalities such as CT and MRI has shown significant potential for improving diagnostic accuracy and patient outcomes. This review has highlighted recent advances:

- (1) Radiomics methods have made significant progress in pancreatic cancer diagnosis, integrating clinical features, employing feature selection techniques, and utilizing various machine learning classifiers such as SVM and XGBoost, with recent advances focusing on application aspects and comparisons between CT and MRI imaging modalities.
- (2) Deep learning approaches have demonstrated potential in automating pancreatic tumor detection and classification by employing CNNs, transformer-based models, and novel deep learning architectures focusing on pancreatic lesions, organ, and tumor segmentation, as well as incorporating auxiliary information to further improve model performance.

Despite the progress made thus far, several challenges remain to be addressed before these methods can be fully integrated into clinical practice. These challenges include the development of fusion models, the establishment of large volume public datasets and standardized benchmarks, the improvement of generalizability across tumor types, domains, and organs, and integrating these functions into CAD systems.

To move the field forward, researchers must address these challenges and refine the current methods to enhance their utility in the clinical setting. Future research should focus on developing more robust, interpretable, and clinically relevant models that can leverage the complementary information from both methods and improve the personalized management of pancreatic cancer patients. The collaboration between experts in AI, medical imaging, pancreatology and oncology will be vital in advancing the field and ultimately improving the early and accurate diagnosis of pancreatic cancer. These efforts, if successful, will contribute to better treatment planning and personalized care, ultimately improving patient outcomes.

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Conflicts of interest

There are no conflicts of interest.

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